

***Amendments to the Claims***

The listing of claims will replace all prior versions, and listings of claims in the application.

1. (previously presented) A composition comprising:

- (a) a virus-like particle;
- (b) at least one immunostimulatory substance; and
- (c) at least one antigen or antigenic determinant;

wherein said at least one antigen or antigenic determinant is bound to said virus-like particle, and wherein said immunostimulatory substance is packaged into said virus-like particle, and wherein said immunostimulatory substance is an immunostimulatory nucleic acid, and wherein said antigen comprises at least one HIV polypeptide.

2. (previously presented) The composition of claim 1, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one nonpeptide covalent bond.

3. (cancelled)

4. (withdrawn-previously presented) The composition of claim 1, wherein said at least one HIV polypeptide is selected from:

- (a) HIV protein subunit p17-GAG;
- (b) HIV protein subunit p24-GAG;
- (c) HIV protein subunit p15-GAG;
- (d) HIV protein subunit Protease;
- (e) HIV protein subunit reverse transcriptase (RT);
- (f) HIV protein subunit Integrase;
- (g) HIV protein subunit Vif;
- (h) HIV protein subunit Vpr;

- (i) HIV protein subunit Vpu;
- (j) HIV protein subunit Tat;
- (k) HIV protein subunit Rev
- (l) HIV protein subunit gp-41-Env;
- (m) HIV protein subunit gp-120-Env;
- (n) HIV protein subunit Nef;
- (o) Nef-protein consensus sequence (SEQ ID NO: 75);
- (p) GAG consensus sequence (SEQ ID NO: 76); and
- (q) any fragment of any of the HIV protein subunits or consensus sequences from (a) to (p).

5. (cancelled)

6. (withdrawn-Previously presented) The composition of claim 1, wherein said at least one HIV polypeptide has the amino acid sequence of Nef-protein consensus sequence (SEQ ID NO: 75), GAG consensus sequence (SEQ ID NO: 76), or a fragment thereof.

7. (withdrawn-Previously presented) The composition of claim 1, wherein said at least one HIV polypeptide comprises an amino acid sequence selected from:

- (a) the amino acid sequence of SEQ ID NO: 77;
- (b) the amino acid sequence of SEQ ID NO: 78;
- (c) the amino acid sequence of SEQ ID NO: 80;
- (d) the amino acid sequence of SEQ ID NO: 81;
- (e) the amino acid sequence of SEQ ID NO: 82;
- (f) the amino acid sequence (SEQ ID NO: 100);
- (g) the amino acid sequence (SEQ ID NO: 102);
- (h) the amino acid sequence (SEQ ID NO: 114);
- (i) the amino acid sequence (SEQ ID NO: 116); and
- (j) any fragment of any of the sequences from (a) to (i).

8. (previously presented) The composition of claim 1, wherein said antigen is a combination of at least two HIV polypeptides, wherein said at least two HIV polypeptides are bound to each other directly or by way of a linking sequence.

9. (withdrawn-previous presented) The composition of claim 8, wherein each of said at least two HIV polypeptides are selected from

- (a) HIV protein subunit p24-GAG;
- (b) HIV protein subunit Nef;
- (c) Nef-protein consensus sequence (SEQ ID NO: 75);
- (d) GAG consensus sequence (SEQ ID NO: 76);
- (e) any fragment of any of the HIV protein subunits or consensus sequences from (a) to (d).

10. (withdrawn-original) The composition of claim 8, wherein said at least two HIV polypeptides are a combination of at least one HIV polypeptide selected from Nef-protein consensus sequence (SEQ ID NO: 75) or a fragment thereof, and of at least one HIV polypeptide selected from GAG-protein consensus sequence (SEQ ID NO: 76) or a fragment thereof.

11. (withdrawn-previous presented) The composition of claim 8, wherein said at least two HIV polypeptides comprise an amino acid sequence selected from:

- (a) the amino acid sequence of SEQ ID NO: 83;
- (b) the amino acid sequence of SEQ ID NO: 84;
- (c) the amino acid sequence of SEQ ID NO: 86;
- (d) any fragment of any of the sequences from (a) to (c).

12. (currently amended) The composition of claim 1 ~~or 8~~, wherein said virus-like particle comprises at least one first attachment site and wherein said antigen or antigenic

determinant further comprises at least one second attachment site being selected from the group consisting of:

- (a) an attachment site not naturally occurring with said antigen or antigenic determinant; and
- (b) an attachment site naturally occurring with said antigen or antigenic determinant;

and wherein said binding of said antigen or antigenic determinant to said virus-like particle is effected through association between said first attachment site and said second attachment site, wherein said antigen or antigenic determinant and said virus-like particle interact through said association to form an ordered and repetitive antigen array.

13. (cancelled)

14. (previously presented) The composition of claim 12, wherein said first attachment site comprises an amino group.

15. (previously presented) The composition of claim 12, wherein said second attachment site comprises a sulphydryl group.

16. (cancelled)

17. (previously presented) The composition of claim 12, wherein said first attachment site is an amino group and said second attachment site is a sulphydryl group.

18. (cancelled)

19. (previously presented) The composition of claim 1, wherein said antigen or antigenic determinant comprises an amino acid sequence selected from:

- (a) the amino acid sequence of SEQ ID NO: 71; and
- (b) the amino acid sequence of SEQ ID NO: 73.

20. (cancelled)

21. (previously presented) The composition of claim 1, wherein said virus-like particle is a recombinant virus-like particle, wherein said virus like particle comprises recombinant proteins selected from the group consisting of:

- (a) recombinant proteins of Hepatitis B virus;
- (b) recombinant proteins of measles virus;
- (c) recombinant proteins of Sindbis virus;
- (d) recombinant proteins of Rotavirus;
- (e) recombinant proteins of Foot-and-Mouth-Disease virus;
- (f) recombinant proteins of Retrovirus;
- (g) recombinant proteins of Norwalk virus;
- (h) recombinant proteins of human Papilloma virus;
- (i) recombinant proteins of BK virus;
- (j) recombinant proteins of bacteriophages;
- (k) recombinant proteins of RNA-phages;
- (l) recombinant proteins of Ty; and
- (m) fragments of any of the recombinant proteins from (a) to (l).

22. (cancelled)

23. (cancelled )

24. (previously presented) The composition of claim 1, wherein said virus-like particle comprises recombinant proteins, or fragments thereof, of a RNA-phage, wherein said RNA-phage is selected from the group consisting of:

- (a) bacteriophage Q $\beta$ ;
- (b) bacteriophage R17;
- (c) bacteriophage fr;
- (d) bacteriophage GA;

- (e) bacteriophage SP;
- (f) bacteriophage MS2;
- (g) bacteriophage M11;
- (h) bacteriophage MX1;
- (i) bacteriophage NL95;
- (j) bacteriophage f2;
- (k) bacteriophage PP7; and
- (l) bacteriophage AP205.

25. (previously presented) The composition of claim 1, wherein said virus-like particle comprises recombinant proteins, or fragments thereof, of bacteriophage Q $\beta$  or bacteriophage AP205.

26. (cancelled)

27. (previously presented) The composition of claim 1, wherein said immunostimulatory nucleic acid is selected from the group consisting of:

- (a) ribonucleic acids;
- (b) deoxyribonucleic acids;
- (c) chimeric nucleic acids; and
- (d) any mixtures of at least one nucleic acid of (a), (b) and/or (c).

28. (cancelled)

29. (cancelled)

30. (previously presented) The composition of claim 1, wherein said immunostimulatory substance is an unmethylated CpG-containing oligonucleotide.

31. (cancelled)

32. (cancelled)

33. (previously presented) The composition of claim 30, wherein said unmethylated CpG-containing oligonucleotide comprises a palindromic sequence.

34. (cancelled)

35. (previously presented) The composition of claim 30, wherein said unmethylated CpG-containing oligonucleotide consists of the sequence GGGGGGGGGGGACGATCGTCGGGGGGGGGG (SEQ ID NO: 41).

36.-41. (cancelled)

42. (previously presented) The composition of claim 33, wherein said palindromic sequence comprises GACGATCGTC (SEQ ID NO: 1).

43.-47. (cancelled)

48. (previously presented) The composition of claim 1, wherein said antigen comprises a cytotoxic T cell epitope, a Th cell epitope or a combination of at least two of said epitopes, wherein said at least two epitopes are bound to each other directly or by way of a linking sequence, and wherein said cytotoxic T cell epitope is a viral or a-tumor cytotoxic T cell epitope.

49.-112.(cancelled)

113. (previously presented) The composition of claim 1, wherein said at least one HIV polypeptide consists of an amino acid sequence selected from:

(a) the amino acid sequence of SEQ ID NO: 77;

- (b) the amino acid sequence of SEQ ID NO: 78;
- (c) the amino acid sequence of SEQ ID NO: 80;
- (d) the amino acid sequence of SEQ ID NO: 81;
- (e) the amino acid sequence of SEQ ID NO: 82;
- (f) the amino acid sequence of SEQ ID NO: 100;
- (g) the amino acid sequence of SEQ ID NO: 102,
- (h) the amino acid sequence of SEQ ID NO: 114;
- (i) the amino acid sequence of SEQ ID NO: 116; and
- (j) any fragment of any of the sequences from (a) to (i).

114. (cancelled)

115. (previously presented) The composition of claim 1, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage Q $\beta$ .

116. (previously presented) The composition of claim 30, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage Q $\beta$ .

117. (previously presented) The composition of claim 35, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage Q $\beta$ .

118. (currently amended) The composition of claim 12, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage Q $\beta$ . wherein said at least one antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond, and wherein said first attachment site ~~comprises~~ is an amino group and wherein said second attachment site is a sulphydryl group.

119. (new) The composition of claim 8, wherein said virus-like particle comprises at least one first attachment site and wherein said antigen or antigenic

determinant further comprises at least one second attachment site being selected from the group consisting of:

- (a) an attachment site not naturally occurring with said antigen or antigenic determinant; and
- (b) an attachment site naturally occurring with said antigen or antigenic determinant;

and wherein said binding of said antigen or antigenic determinant to said virus-like particle is effected through association between said first attachment site and said second attachment site, wherein said antigen or antigenic determinant and said virus-like particle interact through said association to form an ordered and repetitive antigen array, and wherein said at least two HIV polypeptides with said second attachment site comprise an amino acid sequence selected from:

- (i) the amino acid sequence of SEQ ID NO: 72;
- (ii) the amino acid sequence of SEQ ID NO: 85;
- (iii) the amino acid sequence of SEQ ID NO: 87; and
- (iv) a fragment of any one of the sequences from (i) to (iii).

120. (new) The composition of claim 1, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage.

121. (new) The composition of claim 30, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage.

122. (new) The composition of claim 35, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage.

123. (new) The composition of claim 42, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage.

124. (new) The composition of claim 123, wherein said RNA-bacteriophage is RNA-bacteriophage Q $\beta$ .

125. (new) The composition of claim 1, wherein said at least one HIV polypeptide is fused to the virus-like particle.

126. (new) The composition of claim 125, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage.

127. (new) The composition of claim 125, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage AP205.

128. (new) The composition of claim 127, wherein said immunostimulatory substance is an unmethylated CpG-containing oligonucleotide.

129. (new) The composition of claim 127, wherein said unmethylated CpG-containing oligonucleotide consists of the sequence  
**GGGGGGGGGGGACGATCGTCGGGGGGGGG (SEQ ID NO: 41).**

130. (new) The composition of claim 116, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

131. (new) The composition of claim 117, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

132. (new) The composition of claim 121, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

133. (new) The composition of claim 122, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

134. (new) The composition of claim 123, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

135. (new) The composition of claim 124, wherein said antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond.

136. (new) The composition of claim 12, wherein said virus-like particle is a virus-like particle of an RNA-bacteriophage, wherein said at least one antigen or antigenic determinant is bound to said virus-like particle by at least one non-peptide covalent bond, and wherein said first attachment site is an amino group and wherein said second attachment site is a sulphydryl group.

137. (new) The composition of claim 136, wherein said immunostimulatory substance is an unmethylated CpG-containing oligonucleotide.

138. (new) The composition of claim 137, wherein said unmethylated CpG-containing oligonucleotide comprises a palindromic sequence.

139. (new) The composition of claim 137, wherein said unmethylated CpG-containing oligonucleotide consists of the sequence  
GGGGGGGGGGGACGATCGTCGGGGGGGGGG (SEQ ID NO: 41).

140. (new) The composition of claim 138, wherein said palindromic sequence comprises GACGATCGTC (SEQ ID NO: 1).

141. (new) The composition of claim 137, wherein said virus-like particle comprises recombinant proteins, and wherein said recombinant proteins consist of coat proteins consisting of the amino acid sequence of SEQ ID NO:10.

142. (new) The composition of claim 139, wherein said virus-like particle comprises recombinant proteins, and wherein said recombinant proteins consist of coat proteins consisting of the amino acid sequence of SEQ ID NO:10.

143. (new) The composition of claim 140, wherein said virus-like particle comprises recombinant proteins, and wherein said recombinant proteins consist of coat proteins consisting of the amino acid sequence of SEQ ID NO:10.

144. (new) The composition of claim 130, wherein said virus-like particle comprises at least one first attachment site and wherein said antigen or antigenic determinant further comprises at least one second attachment site being selected from the group consisting of:

- (a) an attachment site not naturally occurring with said antigen or antigenic determinant; and
- (b) an attachment site naturally occurring with said antigen or antigenic determinant;

and wherein said binding of said antigen or antigenic determinant to said virus-like particle is effected through association between said first attachment site and said second attachment site, wherein said antigen or antigenic determinant and said virus-like particle interact through said association to form an ordered and repetitive antigen array, and wherein said first attachment site is an amino group and wherein said second attachment site is a sulphydryl group.

145. (new) The composition of claim 131, wherein said virus-like particle comprises at least one first attachment site and wherein said antigen or antigenic determinant further comprises at least one second attachment site being selected from the group consisting of:

- (a) an attachment site not naturally occurring with said antigen or antigenic determinant; and
- (b) an attachment site naturally occurring with said antigen or antigenic determinant;

and wherein said binding of said antigen or antigenic determinant to said virus-like particle is effected through association between said first attachment site and said second attachment site, wherein said antigen or antigenic determinant and said virus-like particle interact through said association to form an ordered and repetitive antigen array, and wherein said first attachment site is an amino group and wherein said second attachment site is a sulphydryl group.

146. (new) The composition of claim 135, wherein said virus-like particle comprises at least one first attachment site and wherein said antigen or antigenic determinant further comprises at least one second attachment site being selected from the group consisting of:

- (a) an attachment site not naturally occurring with said antigen or antigenic determinant; and
- (b) an attachment site naturally occurring with said antigen or antigenic determinant;

and wherein said binding of said antigen or antigenic determinant to said virus-like particle is effected through association between said first attachment site and said second attachment site, wherein said antigen or antigenic determinant and said virus-like particle interact through said association to form an ordered and repetitive antigen array, and wherein said first attachment site is an amino group and wherein said second attachment site is a sulphydryl group.